

Application No.: 10582820
Amdt. dated August 8, 2011
Reply to Office Action of April 6, 2011

Amendments to the Claims:

This listing of claims will replace all prior versions, and listings, of claims in the application:

Listing of Claims:

Claim 1 (currently amended): A method for detecting at least one target nucleic acid comprising:

(a) providing at least one set of first and second target-specific nucleic acids, wherein the at least one set of first and second target-specific nucleic acids each comprise nucleotide sequences complementary to a specific one of the at least one target nucleic acid of interest; wherein each of the at least one set of first and second target-specific nucleic acids are specific only for a selected one of the at least one target nucleic acid; wherein the each first target-specific nucleic acid is bound to a selected first affinity tag and the each second target-specific nucleic acid is bound to a selected second affinity tag, wherein the selected first affinity tag is capable of binding specifically to a molecular motor, wherein the molecular motor ~~consists essentially of~~ is a biological or synthetic molecule capable of induced translational or rotational movements that are capable of detection being detected, wherein the selected second affinity tag is capable of binding specifically to a metal nanorod detection probe ~~consisting essentially of a metal nanorod~~;

(b) contacting the at least one set of first and second target-specific nucleic acids to a sample under conditions whereby the at least one set of first and second target-specific nucleic acids will only hybridize to the at least one target nucleic acid if the at least one target nucleic acid is present in the sample, wherein the target nucleic acid and the first and second target-specific nucleic acids' nucleotide-base-pairing-specific-ligation reaction requires formation of upon hybridization to the target nucleic acid the first and second target-specific nucleic acids are juxtaposed at 5' phosphate and 3' hydroxyl termini of two adjacent target-specific nucleic acids which are hybridized to the complementary target nucleic acid to form a nucleic acid strand that contains a first affinity tag and a second affinity tag at the 5' and 3' ends respectively;

Application No.: 10582820
Amdt. dated August 8, 2011
Reply to Office Action of April 6, 2011

- (c) upon hybridization to the at least one target nucleic acid, ligating the first and second target-specific nucleic acids together;
- (d) ~~binding molecular motor to the first affinity tag and the detection probe to the second affinity tag; binding a series of molecular motors on a solid support either before or after assembly with nucleic acid containing affinity tags on the 5' and 3' ends;~~
- (e) binding at least one of the series of molecular motors to the selected first affinity tag;
- (f) binding the metal nanorod detection probe to the selected second affinity tag of the molecular motor-target nucleotide complex either before or after the series of molecular motors is bound to the solid support;
- ([[e]]) g) inducing translational or rotational movement of at least one of the molecular motors ~~motor~~; and
- ([[f]]) h) detecting translational or rotational movement of the at least one molecular motor ~~coupled to the solid support through the detection probe regularly as indicated by changing color~~ detecting changes in light intensity of at least one wavelength originating from the metal nanorod detection probe, where the at least one wavelength indicates the presence of a specific target nucleic acid in the sample, or by observing the metal nanorod detection probe translationally moving wherein differing selected colors of each of the translationally moving metal nanorod detection probes indicate the presence of a unique corresponding target nucleic acid ~~wherein the motor induced movement of the molecular motor serves to detect the target nucleic acid in the sample, and where observation of ATP-dependent rotation of different colored nanorods indicates the presence of a corresponding target nucleic acid each having its unique probe attachment or different motors causing different specific motor-induced motion so as to allow determination of an assortment of different target nucleic acid(s) is/are present in any given sample.~~

Claim 2 (currently amended): The method of claim 1 wherein the method further comprises generating a plurality of target and probe nucleotide base-pairing specific

Application No.: 10582820
Amdt. dated August 8, 2011
Reply to Office Action of April 6, 2011

ligation products following step (c) using ligation chain reaction.

Claim 3 (currently amended): The method of claim 1 wherein the molecular motor comprises ~~consists essentially of~~ an F1-ATPase.

Claims 4-7 (canceled).

Claim 8 (new): The method of claim 1 wherein said detecting comprises detecting oscillation of intensity of light of only one wavelength.

Claim 9 (new): The method of claim 1 wherein said detecting translational or rotational movement of the at least one molecular motor coupled to the solid support comprises attaching the molecular motor onto a nano-electrode and measuring the micro current change or impedance change produced by rotation.

Claim 10 (new): A method for detecting at least one target nucleic acid comprising:

(a) providing at least one set of first and second target-specific nucleic acids, wherein the at least one set of first and second target-specific nucleic acids each comprise nucleotide sequences complementary to a specific one of the at least one target nucleic acid; wherein each of the at least one set of first and second target-specific nucleic acids are specific only for a selected target nucleic acid; wherein each first target specific nucleic acid is bound to a selected first affinity tag and each second target-specific nucleic acid is bound to a selected second affinity tag, wherein the selected first affinity tag is capable of binding specifically to a molecular motor, wherein the molecular motor is a biological or synthetic molecule capable of induced translational or rotational movement that are capable of being detected, wherein the selected second affinity tag is capable of binding specifically to a metal nanorod detection probe;

(b) contacting the at least one set of first and second target-specific nucleic acids to a sample under conditions whereby the at least one set of first and second

Application No.: 10582820
Amdt. dated August 8, 2011
Reply to Office Action of April 6, 2011

target-specific nucleic acids will only hybridize to the at least one target nucleic acid if the at least one target nucleic acid is present in the sample, wherein the target nucleic acid and the first and second target-specific nucleic acids' nucleotide-base-pairing-specific-ligation reaction requires formation of juxtaposed 5' phosphate and 3' hydroxyl termini of two adjacent target-specific nucleic acids which are hybridized to the complementary target nucleic acid to form a nucleic acid strand that contains a first affinity tag and a second affinity tag at the 5' and 3' ends respectively;

- (c) upon hybridization to the at least one target nucleic acid, ligating the first and second target-specific nucleic acids together;
- (d) binding a series of molecular motors on a solid support either before or after assembly with nucleic acid containing affinity tags on the 5' and 3' ends;
- (e) binding at least one of the series of molecular motors to the selected first affinity tag;
- (f) binding the metal nanorod detection probe to the selected second affinity tag of the molecular motor-target nucleotide complex either before or after the series of molecular motors is bound to the solid support;
- (g) inducing translational or rotational movement of at least one of the molecular motors; and
- (h) microscopically detecting translational or rotational movement of the at least one molecular motor coupled to the solid support as indicated by detecting changes in light intensity of at least one wavelength originating from the metal nanorod detection probe, where the at least one wavelength indicates the presence of a specific target nucleic acid in the sample, or by observing the metal nanorod detection probe translationally moving wherein differing selected colors of each of the translationally moving metal nanorod detection probes indicate the presence of a unique corresponding target nucleic acid in the sample.

Claim 11 (new): The method of claim 10 wherein said microscopically detecting comprises using a microscopy technique selected from the group consisting of dark field microscopy and atomic force microscopy.

Application No.: 10582820
Amdt. dated August 8, 2011
Reply to Office Action of April 6, 2011

Claim 12(new): The method of claim 10 further comprising attaching a fluorescent label on a non-rotating part of the molecular motor before microscopically detecting translation or rotational movement, where the metal nanorod detection probe is a quencher metal nanorod detection probe, and wherein microscopically detecting translational or rotational movement comprises observing rotation through periodic quenching of a fluorescence signal by the quencher metal nanorod detection probe.

Claim 13 (new): The method of claim 1 further comprising attaching a fluorescent label on a non-rotating part of the molecular motor before detecting translation or rotational movement, where the metal nanorod detection probe is a quencher metal nanorod detection probe, and wherein detecting translational or rotational movement comprises observing rotation through periodic quenching of a fluorescence signal by the quencher metal nanorod detection probe.

Claim 14 (new): The method of claim 1 wherein said detecting translational or rotational movement of the at least one molecular motor coupled to the solid support comprises using a detection technique selected from the group consisting of dark field microscopy, single molecule fluorescence resonance energy transfer, fluorescence lifetime anisotropy, atomic force microscopy, single molecule anisotropy measurement, and using a surface plasmon resonance biosensor to measure the surface plasmon resonance change during metallic nanorod rotation.